

# Rabbit Anti-phospho-AKT2 (Ser474) antibody

SL4089R

Product Name:	phospho-AKT2 (Ser474)		
Chinese Name:			
Alias:	AKT2 (phospho S474); p-AKT2 (phospho S474); Akt2(Phospho-Ser474); C AKT; MGC9965; Oncogene AKT1; PKB; PRKBA; Protein Kinase B Alpha; RAC Alpha; RAC; RAC PK Alpha; RAC Serine/Threonine Protein Kinase; vAKT Murine Thymoma Viral Oncogene Homolog 1.		
Organism Species:	Rabbit		
Clonality:	Polyclonal		
React Species:	Human, Mouse, Rat, Chicken, Dog, Pig, Cow, Rabbit, Sheep,		
Applications:	WB=1:500-2000ELISA=1:500-1000IHC-P=1:400-800IHC-F=1:400-800IF=1:100- 500 (Paraffin sections need antigen repair) not yet tested in other applications. optimal dilutions/concentrations should be determined by the end user.		
Molecular weight:	56kDa		
Cellular localization:	The nucleuscytoplasmicThe cell membrane		
Form:	Lyophilized or Liquid		
<b>Concentration:</b>	1mg/ml		
immunogen:	KLH conjugated Synthesised phosphopeptide derived from human AKT2 around the phosphorylation site of Ser474:QF(p-S)YS		
Lsotype:	IgG		
Purification:	affinity purified by Protein A		
Storage Buffer:	0.01M TBS(pH7.4) with 1% BSA, 0.03% Proclin300 and 50% Glycerol.		
Storage:	Store at -20 °C for one year. Avoid repeated freeze/thaw cycles. The lyophilized antibody is stable at room temperature for at least one month and for greater than a year when kept at -20°C. When reconstituted in sterile pH 7.4 0.01M PBS or diluent of antibody the antibody is stable for at least two weeks at 2-4 °C.		
PubMed:	PubMed		
Product Detail:	This gene is a putative oncogene encoding a protein belonging to a subfamily of serine/threonine kinases containing SH2-like (Src homology 2-like) domains. The gene		

was shown to be amplified and overexpressed in 2 of 8 ovarian carcinoma cell lines and 2 of 15 primary ovarian tumors. Overexpression contributes to the malignant phenotype of a subset of human ductal pancreatic cancers. The encoded protein is a general protein kinase capable of phophorylating several known proteins. [provided by RefSeq, Jul 2008]

#### Function:

AKT2 is one of 3 closely related serine/threonine-protein kinases (AKT1, AKT2 and AKT3) called the AKT kinase, and which regulate many processes including metabolism, proliferation, cell survival, growth and angiogenesis. This is mediated through serine and/or threonine phosphorylation of a range of downstream substrates. Over 100 substrate candidates have been reported so far, but for most of them, no isoform specificity has been reported. AKT is responsible of the regulation of glucose uptake by mediating insulin-induced translocation of the SLC2A4/GLUT4 glucose transporter to the cell surface. Phosphorylation of PTPN1 at 'Ser-50' negatively modulates its phosphatase activity preventing dephosphorylation of the insulin receptor and the attenuation of insulin signaling. Phosphorylation of TBC1D4 triggers the binding of this effector to inhibitory 14-3-3 proteins, which is required for insulinstimulated glucose transport. AKT regulates also the storage of glucose in the form of glycogen by phosphorylating GSK3A at 'Ser-21' and GSK3B at 'Ser-9', resulting in inhibition of its kinase activity. Phosphorylation of GSK3 isoforms by AKT is also thought to be one mechanism by which cell proliferation is driven. AKT regulates also cell survival via the phosphorylation of MAP3K5 (apoptosis signal-related kinase). Phosphorylation of 'Ser-83' decreases MAP3K5 kinase activity stimulated by oxidative stress and thereby prevents apoptosis. AKT mediates insulin-stimulated protein synthesis by phosphorylating TSC2 at 'Ser-939' and 'Thr-1462', thereby activating mTORC1 signaling and leading to both phosphorylation of 4E-BP1 and in activation of RPS6KB1. AKT is involved in the phosphorylation of members of the FOXO factors (Forkhead family of transcription factors), leading to binding of 14-3-3 proteins and cytoplasmic localization. In particular, FOXO1 is phosphorylated at 'Thr-24', 'Ser-256' and 'Ser-319'. FOXO3 and FOXO4 are phosphorylated on equivalent sites. AKT has an important role in the regulation of NF-kappa-B-dependent gene transcription and positively regulates the activity of CREB1 (cyclic AMP (cAMP)-response element binding protein). The phosphorylation of CREB1 induces the binding of accessory proteins that are necessary for the transcription of pro-survival genes such as BCL2 and MCL1. AKT phosphorylates 'Ser-454' on ATP citrate lyase (ACLY), thereby potentially regulating ACLY activity and fatty acid synthesis. Activates the 3B isoform of cyclic nucleotide phosphodiesterase (PDE3B) via phosphorylation of 'Ser-273', resulting in reduced cyclic AMP levels and inhibition of lipolysis. Phosphorylates PIKFYVE on 'Ser-318', which results in increased PI(3)P-5 activity. The Rho GTPase-activating protein DLC1 is another substrate and its phosphorylation is implicated in the regulation cell proliferation and cell growth. AKT plays a role as key modulator of the AKT-mTOR signaling pathway controlling the tempo of the process of newborn neurons integration during adult neurogenesis, including correct neuron positioning, dendritic development and synapse formation. Signals downstream of phosphatidylinositol 3-kinase (PI(3)K) to mediate the effects of various growth factors such as platelet-derived growth factor

(PDGF), epidermal growth factor (EGF), insulin and insulin-like growth factor I (IGF-I). AKT mediates the antiapoptotic effects of IGF-I. Essential for the SPATA13-mediated regulation of cell migration and adhesion assembly and disassembly. May be involved in the regulation of the placental development.

One of the few specific substrates of AKT2 identified recently is PITX2. Phosphorylation of PITX2 impairs its association with the CCND1 mRNA-stabilizing complex thus shortening the half-life of CCND1. AKT2 seems also to be the principal isoform responsible of the regulation of glucose uptake. Phosphorylates C2CD5 on 'Ser-197' during insulin-stimulated adipocytes. AKT2 is also specifically involved in skeletal muscle differentiation. Down-regulation by RNA interference reduces the expression of the phosphorylated form of BAD, resulting in the induction of caspase-dependent apoptosis. Phosphorylates CLK2 on 'Thr-343'.

#### Subunit:

Interacts (via PH domain) with MTCP1, TCL1A AND TCL1B. Interacts with CLK2, PBH2 and TRAF6.

### Subcellular Location:

Cytoplasm. Nucleus. Cell membrane; Peripheral membrane protein. Note=Localizes within both nucleus and cytoplasm of proliferative primary myoblasts and mostly within the nucleus of differentiated primary myoblasts. By virtue of the N-terminal PH domain, is recruited to sites of the plasma membrane containing increased PI(3,4,5)P3 or PI(3,4)P2.

# **Tissue Specificity:**

Expressed in all cell types so far analyzed.

# Post-translational modifications:

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	Note=Defects in AKT2 are a cause of susceptibility to breast cancer (BC). AKT2
	promotes metastasis of tumor cells without affecting the latency of tumor development.
	With AKT3, plays also a pivotal role in the biology of glioblastoma.
	Diabetes mellitus, non-insulin-dependent (NIDDM) [MIM:125853]: A multifactorial
•	disorder of glucose homeostasis caused by a lack of sensitivity to the body's own insulin.
	Affected individuals usually have an obese body habitus and manifestations of a
	metabolic syndrome characterized by diabetes, insulin resistance, hypertension and
	hypertriglyceridemia. The disease results in long-term complications that affect the eyes,
	kidneys, nerves, and blood vessels. Note=The disease is caused by mutations affecting
	the gene represented in this entry.
	Hypoinsulinemic hypoglycemia with hemihypertrophy (HIHGHH) [MIM:240900]: A
	disorder characterized by hypoglycemia, low insulin levels, low serum levels of ketone
	bodies and branched-chain amino acids, left-sided hemihypertrophy, neonatal
	macrosomia, reduced consciousness and hypoglycemic seizures. Note=The disease is
	caused by mutations affecting the gene represented in this entry.

# Similarity:

Belongs to the protein kinase superfamily. AGC Ser/Thr protein kinase family. RAC

	subfamily. Contains 1 AGC-kinase C-terminal domain.
	Contains 1 PH domain.
	Contains 1 protein kinase domain.
	SWISS: P31751
	Gene ID: 208
	Database links:
	Entrez Gene: 208 Human
	Entrez Gene: 11652 Mouse
	Entrez Gene: 25233 Rat
	<u>Omim: 164731</u> Human
	SwissProt: P31751 Human
	SwissProt: Q60823 Mouse
	<u>SwissProt: P47197</u> Rat
	<u>Unigene: 631535</u> Human
	<u>Unigene: 177194</u> Mouse
	<u>Unigene: 87066</u> Rat
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	Important Note:
	This product as supplied is intended for research use only, not for use in human, therapeutic or diagnostic applications.









Paraformaldehyde-fixed, paraffin embedded (Rat brain); Antigen retrieval by boiling in sodium citrate buffer (pH6.0) for 15min; Block endogenous peroxidase by 3% hydrogen peroxide for 20 minutes; Blocking buffer (normal goat serum) at 37°C for 30min; Antibody incubation with (phospho-AKT2 (Ser474)) Polyclonal Antibody, Unconjugated (SL4089R) at 1:400 overnight at 4°C, followed by operating according to SP Kit(Rabbit) (sp-0023) instructionsand DAB staining.